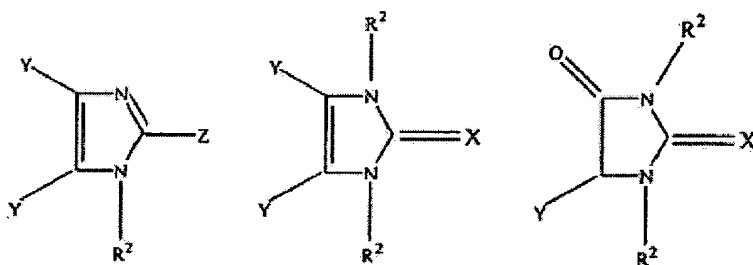


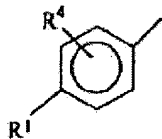
Listing of Claims:

Claims 1-53 (canceled)

54. (previously presented) A method for inhibiting cell adhesion in a mammal which comprises administering to the mammal a pharmaceutical composition comprising a therapeutically effective amount of a compound selected from the group consisting of methimazole derivatives, cyclic thione derivatives, and mixtures thereof, in an amount effective for, inhibition or suppression of cell adhesion, wherein the method is used to treat a cardiovascular disease in a mammal in need of such treatment, wherein the methimazole derivatives have the structural formulae:



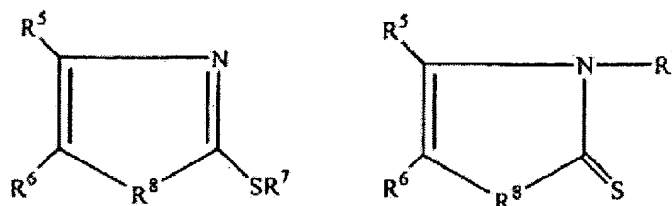
wherein Y is selected from H, Cl -C₄ alkyl C₁ -C₄ substituted alkyl, --NO₂, and the phenyl moiety:



wherein no more than one Y group in said active compound may be the phenyl moiety; R¹ is selected from H, -OH, halogens (F, Cl, Br or I), C₁-C₄ alkyl, C-C₄ substituted alkyl C₁-C₄ ester or C₁-C₄ substituted ester; R² is selected from H, C₁-C₄ alkyl or C₁-C₄ substituted alkyl; R³ is selected from H, C₁-C₄ alkyl, C₁-C₄ substituted alkyl or -CH₂Ph (wherein Ph is phenyl); R⁴ is selected from H, C₁-C₄ alkyl or C₁-C₄ substituted alkyl; X is selected from S or O; Z is selected from -SR³, -OR³, S(O)R³ or C₁-C₄ alkyl; and wherein at least two of the R² and R³ groups on

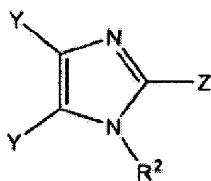
from $-SR^3$, $-OR^3$, $S(O)R^3$ or C_1-C_4 alkyl; and wherein at least two of the R^2 and R^3 groups on said compound are C_1-C_4 alkyl when Y is not a phenyl moiety, and at least one Y is $-NO_2$ when Z is alkyl; together with a pharmaceutically-acceptable carrier; and

wherein the cyclic thione derivatives have the structural formulae:

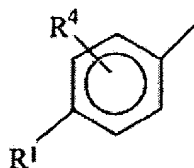


wherein R^5 is selected from CH_3 , CH_3 ; Ph, H; H, and a phenyl moiety; wherein R^6 is selected from CH_3 , CH_3 ; Ph, H; H, and a phenyl moiety; wherein R^7 is selected from H and CH_3 ; and R^8 is selected from O, S, NH, and NCH_3 .

55. (previously presented) The method of claim 54 wherein the compound has a formula:



wherein Y is selected from the group consisting of H, C_1-C_4 alkyl, C_1-C_4 substituted alkyl, $-NO_2$, and the phenyl moiety:

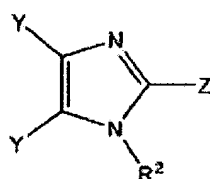


and wherein no more than one Y group in said active compound may be the phenyl moiety; R^1 is selected from the group consisting of H, $-OH$, halogens, C_1-C_4 alkyl, and C_1-C_4

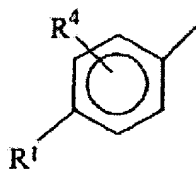
substituted alkyl; R^2 is selected from the group consisting of H, C_1 - C_4 alkyl, C_1 - C_4 substituted alkyl, and a phenyl moiety; R^4 is selected from the group consisting of H, C_1 - C_4 alkyl, and C_1 - C_4 substituted alkyl; and Z is selected from $-SR^3$ and $-OR^3$; R^3 is H; and wherein R^2 in said compound is C_1 - C_4 alkyl when Y is not a phenyl moiety; and a pharmaceutically-acceptable carrier.

56. (withdrawn) A method for the treatment of diseases, disorders, conditions or symptoms mediated by cell adhesion in a mammal which comprises administering to the mammal a pharmaceutical composition comprising methimazole derivatives and/or cyclic thione derivatives are selected from the group consisting of tautomeric methimazole derivatives, non-tautomeric methimazole derivatives, and non-tautomeric cyclic thione derivatives, and combinations thereof.
57. (previously presented) The method according to claim 55, wherein the cardiovascular disease is atherosclerosis.
58. (previously presented) The method according to claim 54, wherein the cell adhesion is VCAM-I mediated.
59. (previously presented) The method according to claim 58, wherein the cell adhesion is mediated by VCAM-I and E-selectin.
60. (previously presented) The method according to claim 59, wherein the cell adhesion is IRF-1 dependent VCAM-I mediated cell adhesion.
61. (previously presented) The method according to claim 54, wherein the cell adhesion is cytokine-induced cell adhesion.
62. (previously presented) The method according to claim 61, wherein the cell adhesion is mediated by VCAM-I and E-selectin.

63. (previously presented) The method according to claim 62, wherein the cell adhesion is IRF-1 dependent VCAM-I mediated cell adhesion.
64. (previously presented) The method according to claim 61, wherein the cytokine is TNF-alpha.
65. (canceled).
66. (canceled)
67. (canceled)
68. (canceled)
69. (withdrawn) The method according to claims 55, 58 or 61, wherein the pharmaceutical composition comprises a safe and effective amount of the tautomeric methimazole derivative:



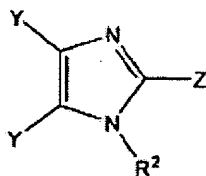
wherein Y is selected from the group consisting of H, C₁-C₄ alkyl, C₁-C₄ substituted alkyl, -NO₂, and the phenyl moiety:



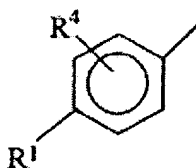
and wherein no more than one Y group in said active compound may be the phenyl moiety; R¹ is selected from the group consisting of H, -OH, halogens, C₁-C₄ alkyl, and C₁-C₄ substituted alkyl; R² is selected from the group consisting of H, C₁-C₄ alkyl, C₁-C₄ substituted

alkyl, and a phenyl moiety; R^3 is H ; R^4 is selected from the group consisting of H, C₁-C₄ alkyl, and C₁-C₄ substituted alkyl; and Z is selected from -SR³ and -OR³1; and wherein R² in said compound is C₁-C₄ alkyl when Y is not a phenyl moiety; and a pharmaceutically-acceptable carrier.

70. (withdrawn) The method according to claims 55, 58 or 61, wherein the pharmaceutical composition comprises a safe and effective amount of the non-tautomeric methimazole derivative



wherein Y is selected from the group consisting of H, C₁-C₄ alkyl, C₁-C₄ substituted alkyl, -NO₂, and the phenyl moiety:

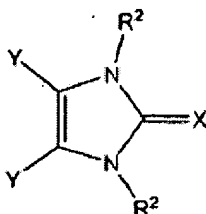


and wherein no more than one Y group in said active compound may be the phenyl moiety; R¹ is selected from the group consisting of H, -OH, halogens, C₁-C₄ alkyl, and C₁-C₄ substituted alkyl; R² is selected from the group consisting of H, C₁-C₄ alkyl, C₁-C₄ substituted alkyl, and a phenyl moiety; R³ is selected from the group consisting of C₁-C₄ alkyl, C₁-C₄ substituted alkyl, and

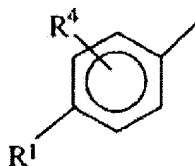
-CH₂Ph; R⁴ is selected from the group consisting of H, C₁-C₄ alkyl, and C₁-C₄ substituted alkyl; and Z is selected from -SR³, S(O)R³, -OR³ and C₁-C₄ alkyl; and wherein at least two of the

R² and R³ groups in said compound are C₁-C₄ alkyl when Y is not a phenyl moiety, and at least one Y is -NO₂ when Z is alkyl; and a pharmaceutically-acceptable carrier.

71. (withdrawn) The method according to claims 55, 58, or 61, wherein the pharmaceutical composition comprises a safe and effective amount of the non-tautomeric cyclic thione derivative



wherein Y is selected from the group consisting of H, C₁-C₄ alkyl, C₁-C₄ substituted alkyl, -NO₂, and the phenyl moiety:



and wherein no more than one Y group in said active compound may be the phenyl moiety; R¹ is selected from the group consisting of H, -OH, halogens, C₁-C₄ alkyl, and C₁-C₄ substituted alkyl; R² is selected from the group consisting of C₁-C₄ alkyl, C₁-C₄ substituted alkyl, and a phenyl moiety; R⁴ is selected from the group consisting of H, C₁-C₄ alkyl, and C₁-C₄ substituted alkyl; X is S; and a pharmaceutically-acceptable carrier.

72. (withdrawn) The method according to claims 69 or 70, wherein Z is SR³ and Y is H.

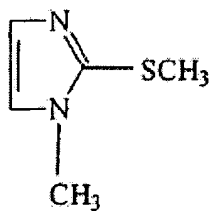
73. (withdrawn) The method according to claim 72, wherein R³ is C₁-C₄ alkyl.

74. (withdrawn) The method according to claim 73, wherein R³ is methyl.

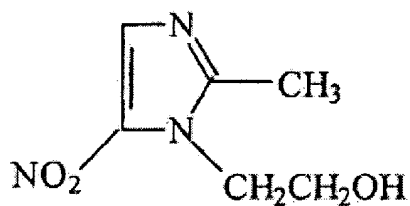
75. (withdrawn) The method according to claim 74, wherein the R² group is methyl.

76. (withdrawn) The method according to claim 74, wherein both R² groups are methyl.

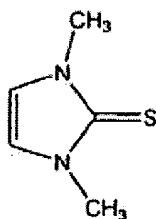
77. (withdrawn) The method according to claim 70, wherein the active compound has the formula



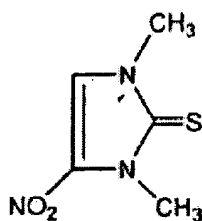
78. (withdrawn) The method according to claim 70, wherein the active compound has the Formula



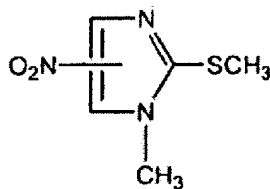
79. (withdrawn) The method according to claim 71, wherein the active compound has the formula:



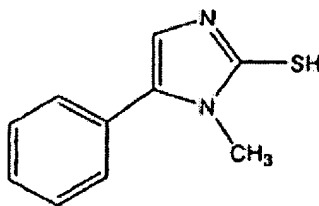
80. (withdrawn) The method according to claim 71, wherein the active compound has the formula:



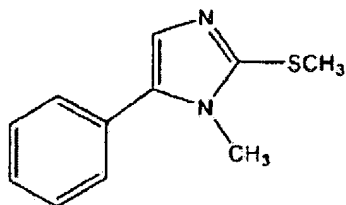
81. (withdrawn) The method according to claim 70, wherein the active compound has the formula:



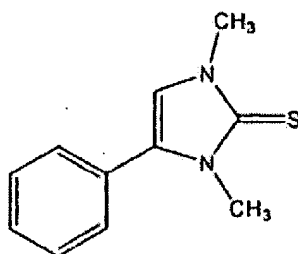
82. (withdrawn) The method according to claims 69 or 70, wherein Z is SR³ and one of the Y groups is the phenyl moiety.
83. (withdrawn) The method according to claim 71, wherein R¹ and R⁴ are H.
84. (withdrawn) The method according to claim 70, wherein Z is SR³ and R³ is a methyl, and one of the Y groups is the phenyl moiety wherein R¹ and R⁴ are H, and the R² group is methyl.
85. (withdrawn) The method according to claim 69 or 70, wherein Z is SR³ and R³ is H, and one of the Y groups is the phenyl moiety wherein R¹ and R⁴ are H, and the R² group is methyl.
86. (withdrawn) The method according to claim 71, wherein one of the Y groups is the phenyl moiety, wherein R¹ and R⁴ are H, and both R² groups are methyl.
87. (withdrawn) The method according to claim 69, wherein the active compound is:



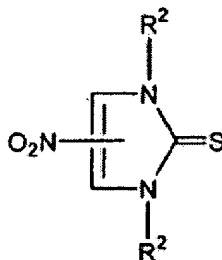
88. (withdrawn) The method according to claim 70, wherein the active compound is



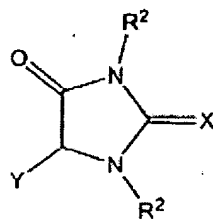
89. (withdrawn) The method according to claim 71, wherein the active compound is



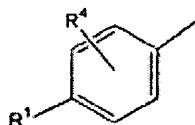
90. (withdrawn) The method according to claims 69, 70, or 71, wherein the pharmaceutical composition is in prodrug form.
91. (withdrawn) The method according to claims 69, 70, or 71, wherein the pharmaceutical composition comprises from about 0.01% to about 25% of the active compound and from about 75% to about 99.99% of the pharmaceutically-acceptable carrier.
92. (withdrawn) The method according to claim 71, wherein the pharmaceutical composition comprises a safe and effective amount of an active compound having the formula:



93. (withdrawn) The method according to claims 55, 58, or 61, wherein the pharmaceutical composition comprises a safe and effective amount of

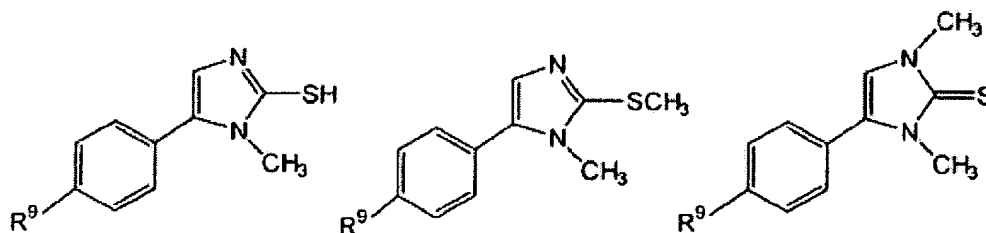


wherein Y is selected from the group consisting of H, C₁-C₄ alkyl, C₁-C₄ substituted alkyl, NO₂ and the phenyl moiety:



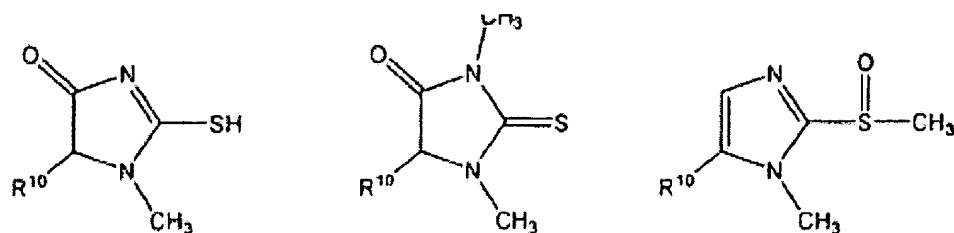
R¹ is selected from the group consisting of H, -OH, halogens, C₁-C₄ alkyl, C₁-C₄ substituted alkyl, C₁-C₄ ester and C₁-C₄ substituted ester; R² is selected from the group consisting of H, C₁-C₄ alkyl and C₁-C₄ substituted alkyl; R⁴ is selected from the group consisting of H, C₁-C₄ alkyl and C₁-C₄ substituted alkyl; X is S; and wherein the R² groups in said compound are C₁-C₄ alkyl when Y is not a phenyl moiety; and a pharmaceutically acceptable carrier.

94. (withdrawn) The method according to claims 55, 58 or 61, wherein the active compound is selected from the group consisting of



wherein R⁹ is selected from the group consisting of -OH, -M and -OOCCH₂M; wherein M is selected from F, Cl, Br and I.

95. (withdrawn) The method according to claims 55, 58, or 61, wherein the active compound is selected from the group consisting of



- wherein R^{10} is selected from the group consisting of H, $-NO_2$, Ph, 4-HOPh and 4-MPh, wherein M is selected from F, Cl, Br and I.
96. (withdrawn) A method of treating a condition involving vascular adhesion of leukocytes, comprising: (a) identifying a subject suspected of having a condition involving aberrant leukocyte adhesion to vascular endothelium; and (b) administering to the subject an amount of a pharmaceutical composition comprising a methimazole derivatives and/or cyclic thione derivatives are selected from the group consisting of tautomeric methimazole derivatives, non-tautomeric methimazole derivatives, and non-tautomeric cyclic thione derivatives, and combinations thereof, sufficient to decrease cell surface expression of at least one of VCAM-I and E-selectin on endothelial cells, thereby reducing adhesion of leukocyte cells to vascular endothelium.
97. (withdrawn) The method according to claim 94, wherein one or more additional active ingredients are combined with the composition of the present invention, either administered separately or in the same pharmaceutical composition, selected from the group comprising (a) VCAM-1 antagonists; (b) steroids; (c) immunosuppressants; (d) antihistamines; (e) non-steroidal anti-asthmatics; (f) non-steroidal antiinflammatory agents (NSAIDs); (g) cyclooxygenase-2 (COX-2) inhibitors; (h) inhibitors of phosphodiesterase type IV (PDE-IV); (i) antagonists of the chemokine receptors; (j)

- cholesterol lowering agents; (k) anti-diabetic agents; (l) preparations of interferon beta; (m) anticholinergic agents; and (n) antibiotics.
98. (withdrawn) A method of screening for compounds capable of ~~preventing~~ inhibiting or suppressing VCAM-1 mediated cell adhesion or preventing, inhibiting or suppressing cell adhesion-associated inflammation in a mammal, comprising:
- (a) contacting the mammal with a test compound;
 - (b) measuring an effect of the test compound on leukocyte adhesion or migration or both; and
 - (c) determining whether the test compound is an inhibitor of the adhesion or migration activities or both of the leukocytes.
99. (withdrawn) The method according to claim 43, wherein the measuring an effect of the test compound comprises one or more of measuring IRF-1 RNA expression levels; measuring IRF-1 protein expression levels; measuring IRF-1 dependent VCAM-1 promoter activation; measuring cytokine -increased IRF-1 RNA expression levels; measuring cytokine -increased IRF-1 protein expression levels; measuring cytokine -increased IRF-1 dependent VCAM-1 promoter activation; measuring TNF-alpha-increased IRF-1 RNA expression levels; measuring TNF-alpha-increased IRF-1 protein expression levels; and measuring TNF-alpha-increased IRF-1 dependent VCAM-1 promoter activation.
100. (withdrawn) The method according to claim 96, wherein the measuring an effect of the test compound comprises one or more of measuring VCAM-1 RNA expression levels; measuring cytokine-increased VCAM-1 protein expression levels; measuring cytokine-increased VCAM-1 promoter activation measuring TNF-alpha-increased VCAM-1 RNA

- expression levels; measuring TNF-alpha increased VCAM-1 protein expression levels;
and measuring TNF-alpha increased VCAM-1
101. (withdrawn) A method of screening for compounds capable of cell adhesion inhibitory activity in VCAM-1 expressing cells, comprising:
- (a) contacting VCAM-1-expressing cells with a test compound;
 - (b) contacting the VCAM-1 -expressing cells to VCAM-1 ligand expressing cells;
 - (c) measuring an effect of the test compound on binding of the VCAM-1-expressing cells to VCAM-1 ligand expressing cells; and
 - (d) determining whether the test compound is an inhibitor of the binding activities of the VCAM-1-expressing cells to VCAM-1 ligand expressing cells.
102. (withdrawn) The method according to claim 99, comprising the further step prior to, concurrently with or subsequently to addition of the test compound, of contacting VCAM-1-expressing cells with a cytokine capable of inducing expression of VCAM-1.
103. (withdrawn) The method according to claim 100, wherein the cytokine is TNF-alpha.
104. (withdrawn) The method according to claim 101, wherein the VCAM-1-expressing cells are selected from the group comprising nonimmune target tissue cells, endothelial cells, and epithelial cells.
105. (withdrawn) The method according to claim 101, wherein the VCAM-1-expressing cells are human aorta endothelial cells (HAEC).
106. (withdrawn) The method according to claim 101, wherein the VCAM-1 ligand expressing cells are leukocytes.
107. (withdrawn) The method according to claim 101, wherein the cells are allowed to remain in contact for at least 30 minutes.